PATENT COOPERATION TRE, .TY

	From the INTERNATIONAL BUREAU	
PCT	То:	
NOTIFICATION OF ELECTION (PCT Rule 61.2) Date of mailing (day/month/year) 14 May 2001 (14.05.01) International application No. PCT/AU00/01143 International filing date (day/month/year) 20 September 2000 (20.09.00) Applicant	Commissioner US Department of Commerce United States Patent and Trademark Office, PCT 2011 South Clark Place Room CP2/5C24 Arlington, VA 22202 ETATS-UNIS D'AMERIQUE in its capacity as elected Office Applicant's or agent's file reference 466267C Priority date (day/month/year) 20 September 1999 (20.09.99)	
HOGG, Philip, John et al		
1. The designated Office is hereby notified of its election made: X In the demand filed with the International Preliminary Examining Authority on: 18 April 2001 (18.04.01)		
The International Bureau of WIPO 34, chemin des Colombettes	Authorized officer	
1211 Geneva 20, Switzerland	G. Bähr	

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International application No.

A.	CLASSIFICATION OF SUBJECT MATTER		
Int. Cl. 7:	C07F 9/20; 9/78; 9/74		
According to	International Patent Classification (IPC) or to bo	th national classification and IPC	
В.	FIELDS SEARCHED		
Minimum docu SEE BELOV	mentation scarched (classification system followed by V	classification symbols)	
Documentation SEE BELOV	searched other than minimum documentation to the ϵ	xtent that such documents are included in t	the fields searched
	base consulted during the international search (name of A): structure, claim 10; arsenoxide; glutathiosenic		
C.	DOCUMENTS CONSIDERED TO BE RELEVAN	T ·	
Category*	Citation of document, with indication, where ag	opropriate, of the relevant passages	Relevant to claim No.
X,Y	Proc.Natl.Acad.Sci, 86, pp 2607-2611 (198 is the primary target for arsenical drugs again pp 2607-2608		1-10; 35, 38,40
x	X Ann. Rev. Microbiol, 46, pp 695-729 (1992) Fairlamb, A "Metabolism and functions of trypanothione in the kinetoplastida" pp 707-708		1-10; 35, 38, 40
X,Y	X,Y Eur J Biochem, 221, pp 285-295 (1994) Cunningham et al "Mechanism of inhibition of trypanothione reductase and glutathione reductase by trivalent organic arsenicals" Table 1, p 288		1-10, 35, 38, 40
х	Further documents are listed in the continuation	on of Box C See patent fami	ily annex
*Special categories of cited documents: "A" document defining the general state of the art which is not considered to be of particular relevance "E" earlier application or patent but published on or after the international filing date "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) "O" document referring to an oral disclosure, use, exhibition or other means "P" document published prior to the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone document of particular relevance, the claimed invention cannot be considered to involve an inventive step when the document is ecombined with one or more other such documents, such combination being obvious to a person skilled in the art document member of the same patent family			
Date of the actual completion of the international search 6 November 2000 Date of mailing of the international search report 1 4 NOV 2000			
AUSTRALIAN	Name and mailing address of the ISA/AU AUSTRALIAN PATENT OFFICE AUSTRALIAN PATENT OFFICE		
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INTERNATIONAL SEARCH REPORT

International application No.

FC1/AU00/01143			
C (Continuation). DOCUMENTS CONSIDERED TO BE RELEVANT			
Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.	
X,Y	Mol Biochem Parasitol, 9, pp 29-35 (1983) Bhargava et al "Effect of arsenical drugs on glutathione metabolism of <i>Litomosoides carinii</i> " pp 29, 31, 34	1-10, 35, 38, 40	
x	Nature, 361(6408), pp 173-6 (1993) Carter et al "Arsenical-resistant trypanosomes lack an unusual adenosine transporter"	1-10, 35, 38, 40	
X	US 3883650 (Friedheim et al.) 13.05.75 Column 3; formula 1	1-10, 35, 38, 40	
Y	Biochimica et Biophysica acta, 628, pp 241-243 (1980) Pisciotto et al	1-10, 35, 38, 40	
X	Chemical Abstracts Registry No 1122-90-3. p-amino phenyl arsenoxide	1-10	
x	Chemical Abstracts Registry No. 637-03-6. Arsenosobenzene	1	
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INTERNATIONAL SEARCH REPORT

International application No.

Box I	Observations where certain claims were found unsearchable (Continuation of item 2 of first sheet)			
This inte	This international search report has not been established in respect of certain claims under Article 17(2)(a) for the following			
1.	Claims Nos:			
	because they relate to subject matter not required to be searched by this Authority, namely:			
2.	Claims Nos: 1-9; 11-14 (in part); 35 (in part); 38 (in part); 40 (in part) because they relate to parts of the international application that do not comply with the prescribed requirements to such an extent that no meaningful international search can be carried out, specifically: Claim 1 defines groups in broad terms (eg, linker groups; cell membrane impermeable pendant group)			
_	which do not have a clear meaning and thus no meaningful search can be carried out. Similarly, its appendages and the claims identified as above insofar as the claims and the appendages include these terms have not been searched.			
3.	Claims Nos:			
	because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a)			
Вох П	Observations where unity of invention is lacking (Continuation of item 3 of first sheet)			
This Inte	ernational Searching Authority found multiple inventions in this international application, as follows:			
1.	As all required additional search fees were timely paid by the applicant, this international search report covers all searchable claims			
2.	As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.			
3.	As only some of the required additional search fees were timely paid by the applicant, this international search report covers only those claims for which fees were paid, specifically claims Nos.:			
4.	No required additional search fees were timely paid by the applicant. Consequently, this international search report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:			
Remark	on Protest			
	No protest accompanied the payment of additional search fees.			



INTERNATIONAL PRELIMINARY EXAMINATION REPORT

(PCT Article 36 and Rule 70)

See Notification of Transmittal of International Preliminary Examination Report (Form PCT/IPEA/416).				
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see				
ent of opinion with regard to novelty, inventive step and industrial applicability				
n the international application				
rations on the international application				
Date of submission of the demand				
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INTERNATIONAL PRELIM. ARY EXAMINATION REPORT

ternational	application	No
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I.	1	Basis of the report		
1.	With regard to the elements of the international application:*			
	\mathbf{x}	X the international application as originally filed.		
		the description,	pages, as originally filed,	
			pages, filed with the demand,	
			pages, received on with the letter of	
		the claims,	pages , as originally filed,	
			pages, as amended (together with any statement) under Article 19.	
			pages, filed with the demand,	
			pages, received on with the letter of	
		the drawings,	pages , as originally filed,	
			pages, filed with the demand,	
			pages, received on with the letter of	
		the sequence listing	g part of the description:	
			pages , as originally filed	
			pages , filed with the demand	
			pages, received on with the letter of	
2.	With regard to the language, all the elements marked above were available or furnished to this Authority in the language in which the international application was filed, unless otherwise indicated under this item. These elements were available or furnished to this Authority in the following language which is: the language of a translation furnished for the purposes of international search (under Rule 23.1(b)).			
	the language of publication of the international application (under Rule 48.3(b)).			
		the language of the and/or 55.3).	translation furnished for the purposes of international preliminary examination (under Rules 55.2	
3.	With prelin	regard to any nucleo ninary examination v	tide and/or amino acid sequence disclosed in the international application, the international was carried out on the basis of the sequence listing:	
	contained in the international application in written form.			
		filed together with	the international application in computer readable form.	
		furnished subseque	ntly to this Authority in written form.	
		furnished subsequen	ntly to this Authority in computer readable form.	
		The statement that to international applications of the statement of the s	the subsequently furnished written sequence listing does not go beyond the disclosure in the ation as filed has been furnished.	
		The statement that the been furnished	he information recorded in computer readable form is identical to the written sequence listing has	
4.		The amendments ha	ive resulted in the cancellation of:	
		the description	on, pages	
		the claims,	Nos.	
		the drawings	, sheets/fig.	
5.		This report has been go beyond the discle	n established as if (some of) the amendments had not been made, since they have been considered to osure as filed, as indicated in the Supplemental Box (Rule 70.2(c)).**	
•	Replace report	ement sheets which ha as "originally filed" ar	ve been furnished to the receiving Office in response to an invitation under Article 14 are referred to in this and are not annexed to this report since they do not contain amendments (Rules 70.16 and 70.17).	
••			ning such amendments must be referred to under item I and annexed to this report	

INTERNATIONAL PRELIMINARY EXAMINATION REPORT

international application No.
PCT/AU00/01143

III.	Non-establishment of opinion with regard to novelty, inventive step and industrial applicability		
 The questions whether the claimed invention appears to be novel, to involve an inventive step (to be not industrially applicable have not been examined in respect of: 			
	the entire international application,		
	X claims Nos: 1-9; 11-14 (in part); 38 (in part) and 40 (in part)		
that no i pendant	hey relate to parts of the International application that do not comply with the prescribed requirements to such an extent aningful search can be carried out. Claim 1 defines groups in broad terms (eg, linker groups; cell membrane impermeable oup) which do not have a clear meaning and thus no meaningful search can be carried out. Similarly, its appendages and identified as above insofar as the claims and the appendages include these terms have not been searched		
:	the said international application, or the said claims Nos. relate to the following subject matter which does not require an international preliminary examination (specify):		
	the description, claims or drawings (indicate particular elements below) or said claims Nos. are so unclear that no meaningful opinion could be formed (specify):		
	the claims, or said claims Nos. are so inadequately supported by the description that no meaningful opinion could be formed. X no international search report has been established for said claim Nos. 1-9; 11-14 (in part) 38 (in part); 40 (in part) insofar as the claims define broad terms as discussed above. However, a cursory look through the literature has identified citations to some of these broad claims as reported in the IPEO in Box V.		

2.	A meaningful international preliminary examination cannot be carried out due to the failure of the nucleotide and/or aming acid sequence listing to comply with the standard provided for in Annex C of the Administrative Instructions:
•	the written form has not been furnished or does not comply with the standard.
	the computer readable form has not been furnished or does not comply with the standard.

v.	Reasoned statement under Article 35(2) with regard to novelty, inventive step or industrial applicability; citations
	and explanations supporting such statement

1.	Statement		
	Novelty (N)	Claims 10, 15-34, 36-37 and 39	YES
		Claims 1-9, 35, 38 and 40-42	NO
	Inventive step (IS)	Claims 15-34, 36-37 and 39	YES
		Claims 1-10, 35, 38 and 40-42	NO
•	Industrial applicability (IA)	Claims 1-42	YES
		Claims	NO

2. Citations and explanations (Rule 70.7)

The following documents identified in the International Search Report have been considered for the purposes of this report:

- D1 Proc Natl Acad Sci (1989)
- D2 Ann Rev Microbiol (1992)
- D3 Eur J Biochem (1994)
- D4 Mol Biochem Parasitol (1983)
- D5 Nature (1993)
- D6 US 3883650
- D7 Biochimica et Biophysica Acta (1980)
- D8 Chemical Abstracts No 112-90-3
- D9 Chemical Abstracts No 637-03-6

Novelty (N) and Inventive Step (IS) Claims 1-10, 35, 38 and 40-42

The broad claims include compounds clearly disclosed and taught in the art. The simple compounds of formula (1) include compounds disclosed in citations D1-D7, eg, arsenoxide derivatives. As noted in Box III of this opinion, claim 1 defines broad terms and no meaningful search can be carried out. However, a cursory look through the literature has identified citations as listed above which clearly disclose and teach the invention as defined in claim 1 and its appendages insofar as these terms form part of the definition of the groups in claim 1.

The applicant submits that claim 1 recites compounds falling within the scope are "substantially cell-membrane impermeable". However, it appears that the broad definition of the terms of claim 1 includes arenoxide compounds and derivatives (eg D6; column 6). It appears that the property regarding "cell-membrane impermeability" is inherent to this broad class of compounds.

.. continued in Supplemental Box

VIII. Certain observations on the international application

The following observations on the clarity of the claims, description, and drawings or on the question whether the claims are fully supported by the description, are made:

Claims 1-9, 11-14, 35 and 40 are not fully supported by the description because of the following broad terms; linker group; spacer group; cell membrane impermeable pendant group.

It would require an undue burden of experimentation on the part of the skilled addressee to determine which groups fall within the scope of the said terms.

Further, it is contended that the scope of the claims is speculative.

INTERNATIONAL PRELIMINARY EXAMINATION REPORT

Supplemental Box

(To be used when the space in any of the preceding boxes is not sufficient)

Continuation of Box V

In regard to the definition of "arsenoxide equivalent", the applicant refers to pages 12-13 of the specification, while there is some indication of the groups which fall within the scope of the term, the definition is not exclusive at pages 12-13. There is reference to typical compounds. In any event, an arsenoxide equivalent is defined as any dithiol reactive species that shows essentially the same affinity towards thiols as -AS=O. It is not clear from your submission as to why the compounds of the prior art as listed above do not fall within the scope of claim 1.

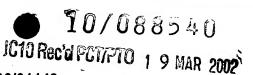
While the specific glutathione derivatives of claim 10 are not disclosed in the art, D9 discloses arsenoxide compounds. These compounds may be further substituted following the teaching in D3.

Moreover the pharmaceutical uses of these compounds is well documented in the art (D1-D7).

he compounds as defined in claim 18-34 appear to be novel and inventive.

Industrial applicability (IA) Claims 1-42

The invention appears to possess industrial applicability.



U.S. NAT'L PHASE OF PCT/AU00/01143 New Claims and Clean Version of Amended Claims

- 6. (Amended) The compound according to claim 1, wherein A is selected from the group consisting of natural, unnatural and synthetic amino acids, hydrophilic amines, peptides, polypeptides, oligosaccharides, detectable groups, and thiol containing proteins, or a combination thereof.
- 9. (Amended) The compound according to claim 1, wherein A is selected from the group consisting of glutathione, glucosamine, cysteinylglycine, cysteic acid, aspartic acid, glutamic acid, lysine, and arginine, and wherein the sulfur atom of each sulfur containing compound may be optionally oxidised to form a sulfoxide or sulfone.
- 10. (Amended) The compound according to claim 1, wherein A is glutathione, and wherein the compound is represented by Formula II:

$$CO_2^ N$$
 H
 S
 L
 $-Y$
 O
 H_2N
 CO_2^-

wherein L comprises any suitable linker and/or spacer group, and Y comprises an arsenoxide or an arsenoxide equivalent.

- 11. (Amended) The compound according to claim 1, wherein p is an integer from 1 to 5.
- 13. (Amended) The compound according to claim 1, wherein L corresponds to (XBX')_nB', and wherein:

n is an integer from 0 to 20,

X is selected from the group consisting of: NR-, S(O)-, -S(O)O-, $-S(O)_2$ -, $-S(O)_2O$ -, -C(O)-, -C(S)-, -C(S)O-, -C(S)S-, $-P(O)(R_1)$ -, $-P(O)(R_1)O$ -, or is absent;

B is selected from C_1 - C_{10} alkylene, C_2 - C_{10} alkenylene, C_2 - C_{10} alkynylene, C_3 - C_{10} cycloalkylene, C_5 - C_{10} cycloalkenylene, C_3 - C_{10} heterocycloalkylene, C_5 - C_{10} heterocycloalkenylene, C_6 - C_{12} arylene, heteroarylene or C_2 - C_{10} acyl;

 $X' \text{ is selected from NR-, $-$O-, $-$S-, $-$S-, $S(O)-, $-$OS(O)O-, $-$OS($

and

B' is C_1 - C_{10} alkylene, C_2 - C_{10} alkenylene, C_2 - C_{10} alkynylene, C_3 - C_{10} cycloalkylene, C_5 - C_{10} heterocycloalkylene, C_5 - C_{10} heterocycloalkenylene, C_6 - C_{12} arylene, heteroarylene or is absent; and wherein

each R is independently selected from hydrogen, C_1 - C_{10} alkyl, C_2 - C_{10} alkenyl, C_2 - C_{10} alkynyl, C_3 - C_{10} cycloalkyl, C_5 - C_{10} cycloalkenyl, C_3 - C_{10} heterocycloalkyl, C_5 - C_{10} heterocycloalkenyl, C_6 - C_{12} aryl, heteroaryl, OR_2 or C_2 - C_{10} acyl;

R' is the same as R or two R' may be taken together with the nitrogen atoms to which they are attached to form a 5 or 6-membered saturated or unsaturated heterocyclic ring;

each R_1 is independently selected from hydrogen, C_1 - C_{10} alkyl, C_2 - C_{10} alkenyl, C_2 - C_{10} alkynyl, C_3 - C_{10} cycloalkyl, C_5 - C_{10} cycloalkenyl, C_3 - C_{10} heterocycloalkyl, C_5 - C_{10} heterocycloalkenyl, C_6 - C_{12} aryl, heteroaryl, halo, OR_2 or NO_2 ;

each R_2 is independently selected from hydrogen, C_1 - C_{10} alkyl, C_2 - C_{10} alkenyl, C_2 - C_{10} alkynyl, C_3 - C_{10} cycloalkyl, C_5 - C_{10} cycloalkenyl, C_3 - C_{10} heterocycloalkyl, C_5 - C_{10} heterocycloalkenyl, C_6 - C_{12} aryl, heteroaryl or $-C(O)R_5$;

each R_5 is independently selected from hydrogen, C_1 - C_{10} alkyl, C_2 - C_{10} alkenyl, C_2 - C_{10} alkynyl, C_3 - C_{10} cycloalkyl, C_5 - C_{10} cycloalkenyl, C_3 - C_{10} heterocycloalkyl, C_5 - C_{10} heterocycloalkyloxy, C_5 - C_{10} alkoxy, C_3 - C_{10} alkenyloxy, C_3 - C_{10} alkynyloxy, C_3 - C_{10} cycloalkyloxy, C_5 - C_{10} cycloalkenyloxy, C_5 - C_{10} heterocycloalkyloxy, C_5 - C_{10} heterocycloalkyloxy, C_5 - C_{10} alkynylthio, C_3 - C_{10} cycloalkylthio, C_5 - C_{10} cycloalkylthio, C_5 - C_{10} cycloalkylthio, C_5 - C_{10} heterocycloalkylthio, C_5 - C_{10} heterocycloalkylthio, C_5 - C_{10} heterocycloalkylthio, C_5 - C_{10} heterocycloalkylthio, C_5 - C_{10} heterocycloalkenylthio, C_6 - C_{12} arylthio, heteroarylthio, C_5 - C_{10} NO₂;

wherein for each instance that B and/or B' is arylene, the substituents directly attached to the respective arylene rings (including arsenoxide or arsenoxide equivalent), may be in a para, meta or ortho relationship, and

wherein each alkylene, alkenylene, alkynylene, cycloalkylene, cycloalkenylene, heterocycloalkylene, heterocycloalkenylene, arylene, heteroarylene and acyl may be independently substituted with hydrogen, C_{1} - C_{10} alkyl, C_{2} - C_{10} alkenyl, C_{2} - C_{10} alkynyl, C_{3} - C_{10} cycloalkyl, C_{5} - C_{10} cycloalkenyl, C_{3} - C_{10} heterocycloalkyl, C_{5} - C_{10} heterocycloalkenyl, C_{6} - C_{12} aryl,

heteroaryl, halo, cyano, cyanate, isocyanate, OR_{2a} , SR_6 , nitro, arsenoxide, $-S(O)R_3$, $-OS(O)R_3$, $-S(O)_2R_3$, $-OS(O)_2R_3$, $-OS(O)_2R_3$, $-OS(O)_2R_4$, $-OP(O)R_4R_4$, $-OP(O)R_4$

wherein R, R₁ and R₅ are as defined above; and

 R_{2a} is selected from hydrogen, C_1 - C_5 alkyl, C_2 - C_5 alkenyl, C_2 - C_5 alkynyl, C_3 - C_{10} cycloalkyl, C_5 - C_{10} cycloalkenyl, C_6 - C_{12} aryl, $-S(O)R_3$, $-S(O)_2R_3$, $-P(O)(R_4)_2$, $N@_2$ or $-C(O)R_5$;

each R_3 is independently selected from hydrogen, C_1 - C_{10} alkyl, C_2 - C_{10} alkenyl, C_2 - C_{10} alkynyl, C_3 - C_{10} cycloalkyl, C_5 - C_{10} cycloalkenyl, C_3 - C_{10} heterocycloalkyl, C_5 - C_{10} heterocycloalkyloxy, C_3 - C_{10} alkoxy, C_3 - C_{10} alkenyloxy, C_3 - C_{10} alkynyloxy, C_3 - C_{10} cycloalkyloxy, C_5 - C_{10} cycloalkenyloxy, C_5 - C_{10} heterocycloalkyloxy, C_5 - C_{10} heterocycloalkyloxy, C_5 - C_{10} alkynylthio, C_3 - C_{10} cycloalkylthio, C_5 - C_{10} alkynylthio, C_5 - C_{10} cycloalkylthio, C_5 - C_{10} heterocycloalkylthio, C_5 - C_{10} heterocycloalkylthio, C_5 - C_{10} heterocycloalkylthio, C_5 - C_{10} heterocycloalkylthio, C_5 - C_{10} heterocycloalkenylthio, C_6 - C_{12} arylthio, heteroarylthio or NO_2 ;

each R_4 is independently selected from hydrogen, C_1 - C_{10} alkyl, C_2 - C_{10} alkenyl, C_2 - C_{10} alkynyl, C_3 - C_{10} cycloalkyl, C_5 - C_{10} cycloalkenyl, C_3 - C_{10} heterocycloalkyl, C_5 - C_{10} heterocycloalkyloxy, C_5 - C_{10} alkoxy, C_3 - C_{10} alkenyloxy, C_3 - C_{10} alkynyloxy, C_3 - C_{10} cycloalkyloxy, C_5 - C_{10} cycloalkenyloxy, C_5 - C_{10} heterocycloalkyloxy, C_5 - C_{10} heterocycloalkenyloxy, C_6 - C_{12} aryloxy, heteroaryloxy, C_1 - C_{10} alkylthio, C_3 - C_{10} alkenylthio, C_3 - C_{10} alkynylthio, C_3 - C_{10} cycloalkylthio, C_5 - C_{10} heterocycloalkylthio, C_6 - C_{12} arylthio, heteroarylthio, halo or NO_2 ;

 R_6 is selected from C_1 - C_{10} alkyl, C_2 - C_{10} alkenyl, C_2 - C_{10} alkynyl, C_3 - C_{10} cycloalkyl, C_5 - C_{10} cycloalkenyl, C_6 - C_{12} aryl, heteroaryl, C_1 - C_{10} alkylthio, C_3 - C_{10} alkenylthio, C_3 - C_{10} alkynylthio, C_3 - C_{10} cycloalkylthio, C_5 - C_{10} cycloalkylthio, C_5 - C_{10} cycloalkylthio, C_5 - C_{10} heterocycloalkylthio, C_5 - C_{10} heterocycloalkenylthio, C_6 - C_{12} arylthio, heteroarylthio, -S(O)R₃, -S(O)₂R₃ or -C(O)R₅.

R" is the same as R or two R" taken together with the N atom to which they are attached may form a saturated, unsaturated or aromatic heterocyclic ring system;

Q is selected from halogen and $-OS(O)_2Q_1$; wherein Q_1 is selected from C_1 - C_4 alkyl, C_1 - C_4 perfluoroalkyl, phenyl, p-methylphenyl; and m is 1 to 5.

18. (Amended) The compound according to claim 1represented by Formula III:

$$R_{3}$$
 R_{3} R_{4} R_{8} R_{8} R_{9} R_{8} R_{9} R_{9

 R_7 to R_{10} are independently selected from the group consisting of: hydrogen, C_1 - C_5 alkyl, C_6 - C_{12} aryl, halogen, hydroxy, amino, nitro, carboxy, C_1 - C_5 alkoxy, $-OS(O)_2R_3$ or $-NHC(O)CH_2Q$ wherein Q is halogen, $-OS(O)_2CH_3$, $-OS(O)_2C_6H_5$ or $-OS(O)_2$ -p tolyl.

- 20. (Amended) The compound according to claim 18, wherein the arsenoxide (-As=O) group is at the 4-position of the phenylene ring.
- 21. (Amended) The compound according to claim 1, wherein the compound is 4-(*N*-(*S*-glutathionylacetyl)amino)phenylarsenoxide (GSAO) and is represented by Formula V:

$$O = \bigvee_{N-H} S \longrightarrow H$$

$$H-N \longrightarrow O$$

$$H_3N^{+} \longrightarrow CO_2$$

$$(V)$$

22. (Amended) The compound according to claim 1, wherein the compound is represented by Formula VI:

wherein Q is any halogen.

23. (Amended) The compound according to claim 1, wherein the compound is represented by Formula VII:

wherein G is selected from the group consisting of: hydrogen, halogen, hydroxy, amino, nitro, carboxy, C_1 - C_5 alkoxy, C_1 - C_5 alkyl and C_6 - C_{12} aryl and -NHC(O)CH₂Q wherein Q is halogen, -OS(O)₂CH₃, -OS(O)₂C₆H₅ or -OS(O)₂-p tolyl.

- 25. (Amended) The compound according to claim 23, wherein G is selected from the group consisting of hydroxy, fluorine, amino, and nitro.
- 26. (Amended) The compound according to claim 23, wherein the activity of the arsenic atom may be modified by the group G, when G and the arsenic atom are in an ortho- or para- relationship to one another.
- 27. (Amended) The compound according to claim 1, wherein the arsenoxide group (-As=O) is replaced by an arsenoxide equivalent.
- 30. (Amended) The compound according to claim 1, which is linked to a detector group.
- 32. (Amended) The compound according to claim 30, wherein the detector group is biotin.
- 35. (Amended) A process for preparing the compound according to claim 1, wherein said process comprises reacting at least one of said substantially cell-membrane impermeable groups (A) with said spacer group L to which is attached at least one arsenoxide or arsenoxide equivalent (Y).
- 37. (Amended) A compound prepared in accordance with the process of claim 35.
- 38. (Amended) A pharmaceutical composition comprising a compound of claim 1, together with a pharmaceutically acceptable carrier, adjuvant and/or diluent.

- 39. (Amended) A process for preparing the pharmaceutical compositioncomprising a compound of claim, wherein said process comprises mixing the compound according to claim 1 with a pharmaceutically acceptable carrier, adjuvant and/or diluent.
- 40. (Amended) A method of treatment and/or prophylaxis of disease in a vertebrate in need of said treatment and/or prophylaxis, wherein said method comprises administering to the vertebrate a therapeutically effective amount of the compound according to claim 1.
- 42. (Amended) The method of claim 40 wherein the disease is selected from the group consisting of angiogenesis-dependent diseases, inflammatory disorders and/or auto-immune diseases, vascular disease and thrombosis, viral infection, and cancer.
- 43. (New) A method of treatment and/or prophylaxis of disease in a vertebrate in need of said treatment and/or prophylaxis, wherein said method comprises administering a therapeutically effective amount of the pharmaceutical composition according to claim 38.
- 44. (New) The compound according to claim 24, wherein G is selected from the group consisting of hydroxy, fluorine, amino, and nitro.